```
L5
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN
     298-14-6 REGISTRY
CN
     Carbonic acid, monopotassium salt (8CI, 9CI)
                                                      (CA INDEX NAME)
OTHER NAMES:
CN
     Armicarb
     Hydrogen potassium carbonate
CN
CN
     K-Lyte
CN
     Kafylox
CN
     Kaligreen
CN
     Monopotassium carbonate
CN
     Potassium acid carbonate
CN
     Potassium bicarbonate
CN
     Potassium bicarbonate (KHCO3)
CN
     Potassium carbonate (KHCO3)
     Potassium hydrogen carbonate (KHCO3)
CN
CN
     Purple K
MF
     C H2 O3 . K
CI
     COM
LC
     STN Files:
                   ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
       BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DDFU, DETHERM*, DIOGENES, DRUGU,
       EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN*, IFICDB,
       IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PDLCOM*, PHAR, PIRA, PROMT, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VTB
          (*File contains numerically searchable property data)
     Other Sources: DSL**, EINECS**, TSCA**
          (**Enter CHEMLIST File for up-to-date regulatory information)
    (463-79-6)
CRN
```

но— с— он

K

3285 REFERENCES IN FILE CA (1957 TO DATE)
35 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
3287 REFERENCES IN FILE CAPLUS (1957 TO DATE)
3 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
=> d

L6 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN 71-52-3 REGISTRY
CN Carbonate, hydrogen (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
```

CN Bicarbonate
CN Bicarbonate (HCO3-)
CN Bicarbonate anion

CN Bicarbonate ion CN Bicarbonate ion (HCO31-)

CN Carbonate (HCO31-)
CN Carbonate ion (HCO31-)
CN Carbonic acid, ion(1-)

CN Carbonic acid, ion(1-)
CN Hydrocarbonate(1-)
CN Hydrogen carbonate

CN Hydrogen carbonate (HCO3-)
CN Hydrogen carbonate anion
CN Hydrogen carbonate ion

CN Hydrogen carbonate ion (HCO3-) CN Monohydrogen carbonate

MF C H O3

CI COM LC STN

STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CBNB, CEN, CHEMINFORMRX, CIN, EMBASE, GMELIN*, IFICDB, IFIPAT, IFIUDB, NIOSHTIC, PIRA, PROMT, SPECINFO, TOXCENTER, TULSA, USPAT2, USPATFULL

(*File contains numerically searchable property data)

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12700 REFERENCES IN FILE CA (1957 TO DATE)
98 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
12710 REFERENCES IN FILE CAPLUS (1957 TO DATE)

```
L2
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN
     63-42-3 REGISTRY
CN
     D-Glucose, 4-O-.beta.-D-galactopyranosyl- (9CI)
                                                           (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN
     Lactose (8CI)
OTHER NAMES:
CN
      (+)-Lactose
CN
     AHL
CN
     Aletobiose
CN
     D-(+)-Lactose
CN
     Fast-flo
     Fast-Flo Lactose
CN
CN
     Galactinum
CN
     Lactin
CN
     Lactin (carbohydrate)
CN
     Lactobiose
CN
     Lactose anhydride
CN
     Lactose anhydrous
CN
     Lactose Fast-flo
CN
     Milk sugar
CN
     Nonpareil 107
CN
     Osmolactan
CN
     Pharmatose 21
     Pharmatose 325M
CN
CN
     Pharmatose 450M
CN
     Saccharum lactin
CN
     Tablettose
CN
     Tablettose 70
CN
     Zeparox EP
AR
     16984-38-6
FS
     STEREOSEARCH
DR
     1336-90-9, 36570-80-6, 73824-63-2, 89466-76-2, 35396-14-6
MF
     C12 H22 O11
CI
     COM
LC
     STN Files:
                   ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
       BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMLIST, CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC,
       PDLCOM*, PIRA, PROMT, RTECS*, SPECINFO, TOXCENTER, TULSA, USPAT2,
       USPATFULL, VETU
          (*File contains numerically searchable property data)
     Other Sources:
                       DSL**, EINECS**, TSCA**
          (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L1ANSWER 1 OF 2 REGISTRY COPYRIGHT 2003 ACS RN 87-78-5 REGISTRY CN Mannitol (8CI, 9CI) (CA INDEX NAME) OTHER NAMES: Mannidex 16700 CN STEREOSEARCH FS DR 133-43-7, 36413-61-3, 5149-40-6 MF C6 H14 O6 CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, DETHERM*, DIOGENES, EMBASE, GMELIN*, HODOC*, IFICDB, IFIPAT, IFIUDB, MEDLINE, NAPRALERT, NIOSHTIC, PDLCOM*, PHARMASEARCH, PIRA, PROMT, RTECS*, TOXCENTER, TULSA, USPAT2, USPATFULL (*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**

(**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

165 REFERENCES IN FILE CA (1957 TO DATE)
12 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
167 REFERENCES IN FILE CAPLUS (1957 TO DATE)
1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

L1 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2003 ACS RN 69-65-8 REGISTRY
CN D-Mannitol (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Cordycepic acid (6CI, 7CI)
CN Mannitol, D- (8CI)
OTHER NAMES:
CN D-(-)-Mannitol

CN Diosmol CN Isotol CN Manicol CNManiton S CN Manna sugar CNMannidex CNMannigen CNMannistol

CN Mannit CN Mannite CN Mannitol CN Mannitolum CN Mannogem 2080 CNMarine Crystal CNOsmitrol CNOsmosal CN Resectisol

STEREOSEARCH

FS

DR 123897-58-5, 75398-80-0, 85085-15-0

MF C6 H14 O6

CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES, DRUGU, EMBASE, GMELIN*, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, NIOSHTIC, PDLCOM*, PHARMASEARCH, PIRA, PROMT, RTECS*, SPECINFO, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VETU

(*File contains numerically searchable property data)
Other Sources: DSL**, EINECS**, TSCA**
 (**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

12451 REFERENCES IN FILE CA (1957 TO DATE)

283 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

12479 REFERENCES IN FILE CAPLUS (1957 TO DATE)

2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
ANSWER 31 OF 294 CA COPYRIGHT 2003 ACS
AN
      136:147485 CA
 TI
      An advantageous carrier solution for vitrifiable concentrations
      of cryoprotectants, and compatible cryoprotectant mixtures
 IN
      Fahy, Gregory M.; Wowk, Brian
PA
      21st Century Medicine, USA
      PCT Int. Appl., 16 pp.
SO
      CODEN: PIXXD2
DT
      Patent
LA
      English
FAN.CNT 4
      PATENT NO.
                          KIND DATE
                                                  APPLICATION NO.
      ------
                          _ _ _ _
                                 -----
                                                  -----
PΙ
      WO 2002009516
                          A2
                                 20020207
                                                  WO 2001-US23853 20010730
                        A3
      WO 2002009516
                                 20020613
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EE, EE, ES, FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD,
               RU, TJ, TM
           RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
               DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
      EP 1311155
                          A2 20030521
                                                  EP 2001-967949 20010730
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
               IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
PRAI US 2000-221691P
                         P
                                 20000731
      WO 2001-US23853
                          W
                                 20010730
      Disclosed herein is a carrier soln. for cryoprotectants that is
AB
      useful for use with cells, tissues, and whole organs and for a variety of
      cryoprotectant solns. and that permits antinucleators to be
      fully effective in vitrification solns., thereby allowing
      vitrification solns. to attain extreme effectiveness, and
      compatible vitrification soln. compns. for use with this carrier
      soln. The carrier soln. comprises lactose and mannitol
      as well as other beneficial ingredients.
IC
      ICM A01N001-02
CC
      9-11 (Biochemical Methods)
ST
     carrier soln vitrifiable concn cryoprotectant compatible
IT
     Animal tissue
     Carriers
     Cell
     Concentration (condition)
     Cryopreservation
     Cryoprotectants
     Kidney
     Mixtures
     Organ, animal
        Solutions
     Vitrification
     Washing
         (advantageous carrier soln. for vitrifiable concns. of
         cryoprotectants, and compatible cryoprotectant mixts.)
IT
     50-99-7, Glucose, biological studies 57-50-1, Sucrose,
     biological studies 63-42-3, Lactose 67-68-5, Dimethyl
     sulfoxide, biological studies 69-65-8, Mannitol
                                                               75-12-7,
     Formamide, biological studies 107-21-1, Ethylene glycol, biological
     studies
                9002-89-5, Polyvinyl alcohol
                                                   9003-20-7D, Poly(vinyl acetate),
     80% hydrolyzed 9003-39-8, Polyvinylpyrrolidone
                                                               9041-07-0, Decaglycerol
     25213-24-5
                   25618-55-7, Polyglycerol 394248-23-8, X 1000
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
```

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L3
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN
     144-55-8 REGISTRY
CN
     Carbonic acid monosodium salt (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     Baking soda
CN
     BI-CF 40E
CN
     BI-H 40E
CN
     Carbonic acid sodium salt (1:1)
CN
     Carbonic acid, monosodium salt
CN
     Cellborn SC-K
CN
     Cellborn SC-P
CN
     Cellmic 266
     DP 35/22
CN
CN
     Extin B
CN
     Meylon
CN
     Monosodium carbonate
CN
     Monosodium hydrogen carbonate
CN
     Soda
CN
     Sodium acid carbonate
CN
     Sodium bicarbonate
CN
     Sodium carbonate (Na(HCO3))
CN
     Sodium hydrogen carbonate
CN
     Sodium monohydrogen carbonate
CN
     Soludal
CN
     Unifine P 4
     196216-68-9, 199723-76-7, 246180-97-2
DR
MF
     C H2 O3 . Na
CI
LC
                   ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOBUSINESS,
     STN Files:
       BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB,
       CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU,
       DETHERM*, DIOGENES, DIPPR*, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2,
       ENCOMPPAT, ENCOMPPAT2, GMELIN*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA,
       MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PDLCOM*, PHARMASEARCH, PIRA, PROMT, RTECS*, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VETU, VTB
          (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
     Other Sources:
          (**Enter CHEMLIST File for up-to-date regulatory information)
CRN
     (463-79-6)
                      Marchiel III 37 53

Marchiel L13 135

Machoiel L25

Vicario
 Na
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```
methacrylate polymer
                       9003-53-6, Polystyrene
                                                 9003-54-7,
 Acrylonitrile-styrene copolymer 9003-56-9
                                               9010-79-1,
 Ethylene-propylene copolymer 9011-14-7, Methyl methacrylate polymer
 24937-78-8, Ethylene-vinyl acetate copolymer 24968-12-5, Polybutylene
 terephthalate
                 25014-41-9, Acrylonitrile polymer
                                                     25038-59-9,
 Polyethylene terephthalate, biological studies
                                                25067-61-2,
 Methacrylonitrile polymer
                            25085-46-5, Ethylene-vinyl acetate-vinyl
 chloride copolymer
                      26062-94-2, Polybutylene terephthalate 120460-98-2,
 Urethane-vinyl chloride copolymer
 RL: BIOL (Biological study)
    (medical goods contg. hemolysis inhibitors and)
 ANSWER 230 OF 294 CA COPYRIGHT 2003 ACS
 102:181076 CA
 Synergistic depression of the freezing temperature in solutions
 of polyhydroxy compounds and antifreeze glycoproteins
 Kerr, William L.; Burcham, Timothy S.; Osuga, David T.; Yeh, Yin; Feeney,
 Robert E.; Caple, Gerald
 Dep. Food Sci. Technol., Univ. California, Davis, CA, 95616, USA
 Cryo-Letters (1985), 6(2), 107-14
                                                                noe &
 CODEN: CRLED9; ISSN: 0143-2044
 Journal
 English
 Mixts. of various polyhydroxy compds. with the low-mol.-wt. fractions of
 antifreeze glycoprotein from Antarctic fish blood showed a greater than
 additive lowering of the freezing temp. Mixts. of polyhydroxy compds.
 with higher-mol.-wt. antifreeze glycoproteins showed a much smaller
 synergistic effect on the freezing temp.
 6-3 (General Biochemistry)
 Section cross-reference(s): 12
 polyhydroxy compd antifreeze glycoprotein synergism; freezing point
 depression polyhydroxy compd glycoprotein; fish glycoprotein polyhydroxy
 compd synergism
 Pagothenia borchgrevinki
    (antifreeze glycoproteins of, f.p. depression by, polyhydroxy compds.
    synergism with)
 Carbohydrates and Sugars, properties
RL: PRP (Properties)
    (f.p. depression response to antifreeze glycoprotein synergism with)
Glycoproteins
RL: BIOL (Biological study)
    (AFGP-4, f.p. depression by, polyhydroxy compds. synergism with)
Glycopeptides
Glycoproteins
RL: BIOL (Biological study)
    (antifreeze, f.p. depression by, polyhydroxy compds. synergism with)
Hydroxy compounds
RL: BIOL (Biological study)
   (poly-, f.p. depression response to antifreeze glycoprotein synergism
   with)
50-70-4, properties 50-99-7, properties
                                          56-81-5, properties
57-50-1, properties
                      59-23-4, properties 63-42-3
69-65-8
          69-79-4
                    87-89-8 87-99-0
                                        488-81-3
RL: PRP (Properties)
   (f.p. depression by antifreeze glycoprotein synergism with)
ANSWER 250 OF 294 CA COPYRIGHT 2003 ACS
98:8173 CA
Water-soluble preparation for making an isotonic nitroglycerin
solution
Muench, Ulrich; Giesselmann, Ewald
Sanol Schwarz-Monheim G.m.b.H., Fed. Rep. Ger.
Ger. Offen., 16 pp.
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L61

AN

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L61

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PA

SO

CODEN: GWXXBX

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Patent
LΑ
     German
FAN.CNT 1
     PATENT NO.
                       KIND DATE
                                            APPLICATION NO.
                                                              DATE
                       ----
                             -----
                                            -----
PΙ
     DE 3109783
                       A1
                             19821007
                                            DE 1981-3109783
                                                              19810313
     DE 3109783
                        C2
                             19870402
     FR 2501675
                        A1
                             19820917
                                            FR 1982-4091
                                                              19820311
     FR 2501675
                        В1
                             19860919
     WO 8203172
                       A1
                             19820930
                                            WO 1982-DE54
                                                              19820312
         W: AT, CH, GB, HU, JP, LU, NL, US
     JP 58500327
                        T2
                             19830303
                                            JP 1982-500862
                                                              19820312
                             19940302
     JP 06015470
                        B4
     GB 2110535
                        A1
                             19830622
                                            GB 1982-32543
                                                              19820312
     GB_2110535
                       B2
                             19850206
     (US 4481220)
                       A
                             19841106
                                            US 1982-438887
                                                              19820929
     AT 8802240
                       Α
                             19890315
                                            AT 1988-2240
                                                              19880913
     AT 389050
                       В
                             19891010
PRAI DE 1981-3109783
                             19810313
     AT 1982-9015
                             19820312
     WO 1982-DE54
                             19820312
AB
     A prepn. that can be dild. with H2O to give an isotonic infusion or
     injection soln. contg. 1 mg/mL nitroglycerin [55-63-0] contains nitroglycerin and a solubilizer in a ratio of 0.5:1 to 2:1 and a solid
     nitroglycerin carrier capable of controlling isotonicity. Thus, 1.6 kg
     nitroglycerin was dissolved in 8-10 L Et20 and mixed with 1.6 kg
     1,2-propylene glycol [57-55-6]. The soln. was mixed with 76.8
     kg glucose [50-99-7] with aeration until the Et2O evapd. and
     the soln. was homogeneous. The soln., 11.550 kg, was
     added gradually to H2O at 70.degree. until dissolved, cooled, and dild.
     with H2O (total of 220 L). The soln. was placed in ampuls or
     bottles for injection.
IC
     A61K031-21
CC
     63-6 (Pharmaceuticals)
ST
     nitroglycerin solubilizer injection; propylene glycol nitroglycerin
     injection
IT
     Solubilizers
        (for nitroglycerin injections)
ΙT
     RL: BIOL (Biological study)
        (injections, solubilizers and carriers for)
ŦΤ
     50-70-4, biological studies 50-99-7, biological studies
     57-48-7, biological studies 63-42-3 69-65-8
     7647-14-5, biological studies
     RL: BIOL (Biological study)
        (nitroglycerin injections contg., for controlling isotonicity)
TT
     51-79-6 97-64-3
                         100-79-8
                                     107-88-0
                                                111-55-7
                                                          111-96-6
                                                                       112-60-7
                                       542-59-6 1187-03-7 1569-01-3
     123-80-8 126-33-0
                          127-19-5
     4128-76-1
               5422-34-4
                             5464-28-8 9003-11-6
                                                     9004-32-4 11111-34-5
     19354-27-9
                  24567-27-9
                               25322-68-3
                                            29387-84-6
                                                          53778-73-7
     83931-54-8
     RL: BIOL (Biological study)
        (nitroglycerin solubilization by, for injections)
IT
     50-21-5, properties
                           56-81-5, properties
                                                 57-13-6, biological studies
     57-55-6, properties
                           68-12-2, properties
     RL: PRP (Properties)
        (nitroglycerin solubilization by, for injections)
    ANSWER 270 OF 294 CA COPYRIGHT 2003 ACS
L61
AN
     85:18011 CA
ΤI
    Lyophilization of hemoglobin solutions. Study of protector
     compounds capable of preventing the formation of methemoglobin
ΑIJ
    Labrude, P.; Vigneron, C.; Streiff, F.
    Cent. Reg. Transfus. Sang. Nancy-Brabois, Vandoeuvre, Fr.
CS
```

DT

```
Journal de Pharmacie de Belgique (1976), 31(2), 191-8
      CODEN: JPBEAJ; ISSN: 0047-2166
DT
     Journal
LA
     French
AB
     The protection of Hb solns. against oxidn. to methemoglobin
     during lyophilization by plasma expanders, macromols., sugars, glycerol,
     and THAM was studied. The most active mols. were the sugars and THAM
     which almost entirely prevented the formation of methemoglobin at a concn.
     of 1.25%.
     13-5 (Mammalian Biochemistry)
     Hb oxidn lyophilization preservative; methemoglobin formation
     lyophilization preservative
IT
     Blood substitutes
     Albumins
     Sugars, biological studies
     RL: BIOL (Biological study)
         (Hb oxidn. response to, in lyophilization)
IT
     Methemoglobins
     RL: FORM (Formation, nonpreparative)
         (formation of, in lyophilization, prevention of)
IT
     Freeze drying
         (of Hb, oxidn. prevention in)
IT
     Hemoglobins
     RL: RCT (Reactant); RACT (Reactant or reagent)
         (oxidn. of, in lyophilization, prevention of)
IT
     50-70-4 50-99-7, biological studies
                                            56-81-5, biological
     studies
               57-48-7, biological studies
                                              57-50-1, biological studies
     59-23-4, biological studies 63-42-3 69-65-8
                                                    77-86-1
     3458-28-4
                 8057-73-6
                             9002-89-5
                                          9003-39-8
                                                      9004-54-0, biological
     studies
               39290-10-3
                             54847-63-1
                                          66455-30-9
     RL: BIOL (Biological study)
        (Hb oxidn. response to, in lyophilization)
L61 ANSWER 272 OF 294 CA COPYRIGHT 2003 ACS
AN
     83:197730 CA
ΤI
     Comparative testing of titrable acidity degree and pH value of infusion
     solutions
ΑU
     Horvath, Klara; Regos, Erika; Varga, Sarolta
CS
     Inst. Serobacteriol. Prod. Res. "HUMAN", Godollo, Hung.
     Annales Immunologiae Hungaricae (1973), 17, 265-8
SO
     CODEN: AIMHA3; ISSN: 0570-1708
DT
     Journal
LA
     English
AB
     The titrable acidity degree (i.e. the hydrogen ion reserve possessed by
     the nondissociated species of weak acids) of various infusion
     solns. was reported and compared with the pH of the same
     solns. Since in the case of weak acids pH does not characterize
     unequivocally the acidic character of the soln. it was suggested
     that the labels of infusion solns. also contain the value of
     titrable acidity.
CC
     63-5 (Pharmaceuticals)
     infusion soln titrable acidity
st
IT
     Ringer's solution
     RL: BIOL (Biological study)
        (acetate infusion soln., titrable acidity degree of, pH in
        relation to)
IT
     Pharmaceuticals
        (infusion solns., titrable acidity degree of, pH in relation
TΤ
     Ringer's solution
     RL: BIOL (Biological study)
        (lactate infusion soln., titrable acidity degree of, pH in
        relation to)
IT
    Ringer's solution
```

RL: BIOL (Biological study)
 (titrable acidity degree of, pH in relation to)

IT 9004-54-0, biological studies
RL: BIOL (Biological study)
 (infusion glucose soln., titrable acidity degree of, pH in relation to)

IT 50-99-7, biological studies 57-48-7, biological studies
63-42-3 69-65-8 9004-54-0, biological studies
57455-70-6 57455-71-7
RL: BIOL (Biological study)
 (infusion soln., titrable acidity degree of, pH in relation

=>

(Uses) (advantageous carrier solm. for vitrifiable concns. of cryoprotectants, and compatible cryoprotectant mixts.) ANSWER 65 OF 294 CA COPYRIGHT 2003 ACS 133:109998 CA AN Beta-interferon lozenge and its preparing method ΤI IN Cao, Xuetao; Ju, Dianwen; Tao, Qun PA Huachen Biological Technology Inst., Shanghai, Peop. Rep./China Faming Zhuanli Shenqing Gongkai Shuomingshu, 13 pp. SO CODEN: CNXXEV DT Patent LAChinese FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE ---------PRAI CN 1998-105383 CN 1227124 19990901 CN 1998-105383 19980225 19980225 The beta interferon lozenge is composed of 1-10 /kIU beta-interferon and medicinal adjuvant. The medicinal adjuvant is selected from one or more of human serum albumin, bovine serum protein, polyethylene glycol, mannitol, lactose, glucose, starch, Mg stearate, and dextrin. The beta-interferon lozenge may contain 100-1,000 IU alpha-interferon and/or 1-10 kIU interleukin-2. The beta interferon lozenge is prepd. by mixing medicinal adjuvant, sieving, drying to obtain blank granule; spraying beta-interferon soln. in the blank granule; drying, and tableting. The lozenge is used for treatment of virus infection and/or tumor. IC ICM A61K038-21 ICS A61K009-20 CC 63-6 (Pharmaceuticals) Section cross-reference(s): 15 STbeta interferon lozenge prepn IT Proteins, general, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (blood; beta-interferon lozenge and its prepg. method) Drug delivery systems (lozenges; beta-interferon lozenge and its prepq. method) TT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (.alpha.; .beta.-interferón lozenge and its prepg. method) IT Antitumor agents Antiviral agents (.beta.-interferon lozenge and its prepg. method) Albumins, biological studies IT Interleukin 2 Polyoxyalkylenes, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (.beta.-interferon lozenge and its prepg. method) ΙT Interferons RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (.beta.; .beta.-interferon lozenge and its prepg. method) 9004-53-9, Dextrin /9005-25-8, Starch, biological studies IT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (beta-interferon/lozenge and its prepg. method) IT 50-99-7, Glucose, biological studies 63-42-3, Lactose

L61 ANSWER 191 OF 294 CA COPYRIGHT 2003 ACS

AN 112:62604 CA

Polyethylene glycol

TI Monocarboxylic acid esters as blood preservatives

69-65-8, Mannitol / 557-04-0, Magnesium stearate

(.beta.-interferon lozenge and its prepq. method)

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

25322-68-3,

```
IN
      Nagai, Hiroshi; Kubota, Yoshiki; Tamura, Yoko; Kimura, Akio
PA
      Terumo Corp., Japan; Kao Corp.
SO
      Jpn. Kokai Tokkyo Koho, 8 pp.
      CODEN: JKXXAF
DT
      Patent
LΑ
     Japanese
FAN.CNT 1
                       KIND DATE
      PATENT NO.
                                           APPLICATION NO.
                       ----
                                             -----
ΡI
     JP 01106826
                       A2
                             19890424
                                             JP 1987-263695
                                                               ⁄19871021
PRAI JP 1987-263695
                             19871021
os
     MARPAT 112:62604
     A preservative for blood contains an antihemolytic agent, RCO2R1 (R and R1
AB
     = C .gtoreq.3 hydrocarbyl; the sum of C in R and P1 is 11-30), together
     with other preservatives such as Na citrate, citric acid, glucose,
     NaH2PO4, adenine, NaCl, mannitol, etc. Thus, Tween-80 was dissolved (600
      .mu.g/mL) in a soln. contg. NaCl 140, adenine /1.25, and glucose
     50 mM, and 12 mM iso-Pr isolaurate was added. This emulsion (1.0 mL) was
     added as a preservative to 2.0 mL human erythrocyte conc. (with hematocrit
     value 70%) and stored for 5 wk at 4.degree.'.
IC
     ICM A61K035-14
CC
     63-3 (Pharmaceuticals)
                                                                          25 5 m 1/m 1
ST
     blood preservative antihemolytic carboxylate
TT
     Blood preservatives
         (contg. monocarboxylic acid esters, as antihemolytic agents)
IT
     Carboxylic acids, esters
     RL: BIOL (Biological study)
         (esters, blood preservatives contg., as antihemolytic agents)
     50-70-4, Sorbitol, biological studies 50-99-7, D-Glucose, biological studies 57-50-1, Sucrose, biological studies 63-42-3, Lactose 68-04-2, Sodium citrate 69-65-8, Mannitol 69-79-4, Maltose 73-24-5, Adenine, biological studies 77-92-9, Citric acid,
TT
     biological studies 585-88-6, Maltitol
                                                 7558-80-7, Monosodium phosphate
     7647-14-5, Sodium chloride, biological studies
     RL: BIOL (Biological study) /
         (blood preservatives contg. antihemolytic carboxylic esters and)
     112-11-8, Isopropyl oleate/ 7425-14-1, 2-Ethylhexyl 2-ethylhexanoate
IT
     81897-25-8, 2-Ethylhexyl isostearate 120470-79-3, Isopropyl isolaurate
     RL: BIOL (Biological study)
         (blood preservatives/contg., as antihemolytic agent)
L61 ANSWER 197 OF 294 CA COPYRIGHT 2003 ACS
ΑN
ΤI
     Hemolysis inhibitors and medical resin compositions, medical goods, and
     blood-preserving fluids containing the same
     Nagai, Hirofumi; Kubota, Yoshinori; Tamura, Yoko; Sado, Mineo; Kora,
IN
     Shinichi; Kimura, Akio; Suzue, Shigetoshi; Miyamoto, Norioki
PA
     Kao Corp., Japan; Terumo Corp.
so
     PCT Int. Appl., 163 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     Japanese
FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                           APPLICATION NO.
                                                              DATE
                     ----
PΤ
     WO 8803027
                      A1 19880505
                                            WO 1987-JP810
                                                              19871022
         W: AU, US
         RW: BE, DE, FR, GB, IT, NL, SE
     JP 63104916 A2 19880510
                                            JP 1986-249552
                                                              19861022
     JP 04015203
                      B4 19920317
     AU 8781050
                      A1 19880525
                                            AU 1987-81050
                                                              19871022
     AU 623071
                     B2 19920507
   ES 2005457
                     A6 19890301
                                           ES 1987-3293
                                                              19871022
                 A1 19891004
```

EP 1987-906934

19871022

EP 334956

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R: BE, DE, FR, GB, IT, NL, SE
PRAI JP 1986-249552
                            19861022
     WO 1987-JP810
                            19871022
AB
     Hemolysis inhibitors contg. carboxylic acid ester polymers, ether compds.,
     and/or monocarboxylic ester are used to manuf. clin. resin compns.,
     medical goods and blood preservers. Iso-Pr isotridecyl maleate (I) was
     dispersed in a mixt. of polyoxyethylene monooleate-saline (2000 .mu.g/mL)
     to make a 4 mM emulsion. This emulsion was added to a fluid contg. human
     erythrocytes having 70% hematocrit value such that the concn. of I was 0.4
         The fluid kept at 4.degree. for 4 wk showed the stability of
     erythrocytes. The stability of erythrocytes was also demonstrated by
     keeping blood in a poly(vinyl chloride) bag which consisted of poly(vinyl
     chloride) 100, didecyl phthalate 30, iso-Pr decyl maleate/20, epoxylated
     soybean oil 10, and a Ca-Zn type stabilizer 0.1 parts by wt.
IC
     ICM A61K035-14
     ICS A61L031-00; A61M005-00; A01N001-02
CC
     63-7 (Pharmaceuticals)
     hemolysis inhibitor medical goods; blood preservative hemolysis inhibitor
ST
IT
     Glycerides, biological studies
     RL: DEV (Device component use); USES (Uses)
        (as hemolysis inhibitors, medical goods contg.)
IT
     Blood preservation
        (carboxylic acid esters and ethers in, as hemolysis inhibitors)
ΙT
     Medical goods
        (hemolysis inhibitor-contg.)
IT
     Hemolysis
        (inhibitors, carboxylic acid derivs. as)
IT
     Acrylic polymers, biological studies
     Polycarbonates, biological studies
     Polyesters, biological studies
     Urethane polymers, biological studies
     RL: BIOL (Biological study)
        (medical goods contg. hemolysis inhibitors and)
ΙT
     Erythrocyte
        (stabilization of, by carboxylic acid derivs.)
     50-70-4, D-Glucitol, biological studies 50-99-7, D-Glucose,
     biological studies 57-50-1, biological studies 63-42-3,
                                                   73-24-5,
             68-04-2 69-65-8, Mannitol 69-79-4
     Adenine, biological studies
                                 77-92-9, Citric acid, biological studies
     585-88-6, Maltitol
                          7558-80-7, Monosodium phosphate 7647-14-5, Sodium
     chloride, biological studies
     RL: BIOL (Biological study)
        (as blood preservative soln. contg.)
TΤ
     120486-11-5
     RL: BIOL (Biological study)
        (as hemolysis inhibitor)
IT
     112-11-8, Isopropyl oleate
                                 7425-14-1, 2-Ethylhexyl 2-ethylhexanoate
     59068-03-0, Glycerin 1,3-bis(2-ethylhexyl) ether 81897-25-8
     93120-93-5, Glycerin 1-butyl\frac{1}{7}3-isostearyl ether
                                                      120422-95-9
                  120470-78-2 /120470-79-3
     120422-96-0
                                               120573-91-3
    RL: DEV (Device component use); USES (Uses)
        (as hemolysis inhibitor, medical goods contg.)
IT
    97-65-4D, esters
                      110-15-6D, Butanedioic acid, esters
                                                              110-16-7D, Maleic
                  110-94-1D, Glutaric acid, esters 111-20-6D, Sebacic acid,
    acid, esters
             124-04-9D, Hexanedioic acid, esters 143-07-7D, Dodecanoic acid,
    esters
             328-42-7D, Oxalacetic acid, esters
    RL: DEV (Device component use); USES (Uses)
        (as hemolysis inhibitors, medical goods contg.)
IT
    84-77-5, Didecyl phthalate '89-04-3, Trioctyl trimellitate
    RL: BIOL (Biological study)
        (as plasticizer, medical goods resin compns. contg.)
    9002-86-2, Poly(vinyl chloride) 9002-88-4, Polyethylene
ΙT
                    9003-21-8, Methyl acrylate polymer
    Polypropylene
                                                         9003-22-9, Vinyl
    acetate-vinyl chloride copolymer 9003-32-1 9003-42-3, Ethyl
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ANSWER 24 OF 39 WPIDS (C) 2003 THOMSON DERWENT
     1998-076784 [07]
AN
                        WPIDS
CR
     1998-593980 [50]
DNC
     C1998-025629
TΙ
     Solution for preservation of biological materials - comprises
     two neutral solutes, especially raffinose and tri methylamine oxide.
DC
     B04 D16 D22 E19 E37 P31
IN
     FERGUSON, A B; WIGGINS, P M; WATSON, J D
      (BIOS-N) BIOSTORE NEW ZEALAND LTD; (BIOS-N) BIOSTORE NZ LTD; (BIOS-N)
PA
     BIOSTORE NEW ZEALAND
CYC
     73...
     (WO 9747192)
PΙ
                   A1 19971218 (199807) * EN
                                               53p
        RW: AT BE CH DE DK EA ES FI FR GB GR IE IT KE LS LU MC MW NL OA PT SD
            SE SZ UG
         W: AL AM AT AU AZ BB BG BR BY CA CH CN CZ DE DK EE ES FI GB GE HU IL
            IS JP KE KG KP KR KZ LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL
            PT RO RU SD SE SG SI SK TJ TM TR TT UA UG UZ VN
     AU 9661412
                   A 19980107 (199820)
     ZA 9710452
                   Α
                      19980826 (199840)#
                                               79p
     US 5879875
                   Α
                      19990309 (199917)#
     NZ 333030
                   Α
                      20000526 (200033)
     EP 1018866
                   A1 20000719 (200036)
         R: CH DE FR GB LI SE
     JP 2000512625 W 20000926 (200051)
                                               50p
     AU 725247
                   B 20001012 (200055)
     AU 2001010037 A 20010315 (200121)#
     AU 742402
                   B 20020103 (200209)#
     EP 1018866
                   B1 20021204 (200303) EN
         R: CH DE FR GB LI SE
     DE 69625254
                   E 20030116 (200313)
ADT WO 9747192 A1 WO 1996-NZ57 19960614; AU 9661412 A AU 1996-61412 19960614,
     WO 1996-NZ57 19960614; ZA 9710452 A ZA 1997-10452 19971120; US 5879875 A
     US 1996-662244 19960614; NZ 333030 A NZ 1996-333030 19960614, WO 1996-NZ57
     19960614; EP 1018866 A1 EP 1996-918938 19960614, WO 1996-NZ57 19960614; JP
     2000512625 W WO 1996-NZ57 19960614, JP 1997-541244 19960614; AU 725247 B
     AU 1996-61412 19960614, WO 1996-NZ57 19960614; AU 2001010037 A Div ex AU
     1996-61412 19960614, AU 2001-10037 20010103; AU 742402 B Div ex AU
     1996-61412 19960614, AU 2001-10037 20010103; EP 1018866 B1 EP 1996-918938
     19960614, WO 1996-NZ57 19960614; DE 69625254 E DE 1996-625254 19960614. EP
     1996-918938 19960614, WO 1996-NZ57 19960614
FDT AU 9661412 A Based on WO 9747192; NZ 333030 A Based on WO 9747192; EP
     1018866 A1 Based on WO 9747192; JP 2000512625 W Based on WO 9747192; AU
     725247 B Previous Publ. AU 9661412, Based on WO 9747192; AU 2001010037 A
     Div ex AU 725247; AU 742402 B Previous Publ. AU 200110037, Div ex AU
     725247; EP 1018866 B1 Based on WO 9747192; DE 69625254 E Based on EP
     1018866, Based on WO 9747192
PRAI WO 1996-NZ57
                      19960614; ZA 1997-10452
                                                 19971120; US 1996-662244
     19960614; AU 2001-10037
                                20010103
AB
          9747192 A UPAB: 20030224
     The following are claimed: (1) a solution for preservation of
     biological materials: (A) comprising: (a) a first neutral solute with a
     molecular weight of at least 335 and a solubility in water of at least 0.3
     M, and (b) a second neutral solute with a molecular weight < 200 and with
     both hydrophilic and hydrophobic moieties; or (B) which is isotonic with
     the biological materials and is free of univalent oxyanions and iodide,
     and (2) preservation of biological materials by: (a) pretreating the
    biological material with a solution which includes sodium
    butyrate, and (b) contacting the biological material with a preservative
     solution.
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The **solution** (A) is free of univalent oxyanions and iodide, and also comprises at least 1 ion from a protein-stabilising end of the Hofmeister series. The first solute is selected from disaccharides and trisaccharides, especially raffinose, trehalose, sucrose, **lactose**

and their analogues. The second neutral solute is selected from trimethylamine oxide, betaine, taurine, sarcosine, glucose, mannose, fructose, ribose, galactose, sorbitol, mannitol, inositol and their analogues. The solution may also comprise sodium sulphate. It may also comprise calcium, which is present as calcium sulphate at a concentration of 1.5-2.0 mM. The solution is in a concentrated form, especially in the form of a solid. Components (a) and (b) are typically present at a ratio of 1.4-1.8:1.

USE - The **solutions** may be used for preservation of materials such as **organs**, **tissues** and **cells** from mammals, marine organisms and plants. They may be used e.g. in treatment of leukaemia. In this case, bone marrow is removed from a patient and contacted with the **solution** for at least 3 days, in order to purge the bone marrow of leukaemic **cells**. The purged bone marrow is then returned to the patient.

ADVANTAGE - The **solutions** are of low toxicity, resulting in fewer side effects when biological materials, such as transplant **organs**, are returned to a patient.

Dwq.0/20

```
L69 ANSWER 5 OF 39 WPIDS (C) 2003 THOMSON DERWENT
AN
     2002-479639 [51]
                        WPIDS
    C2002-136479
DNC
     Vitrification of natural or engineered tissue or organ
TI
     other than blood vessel involves immersing tissue in
     cryoprotectant solutions with increasing concentrations and
     cooling to below glass transition temperature.
     A96 D22 E19 G04 J07
DC
     BROCKBANK, K G M; KHIRABADI, B S; SONG, Y C
IN
PΑ
     (ORGA-N) ORGAN RECOVERY SYSTEMS
CYC
    94
     WO 2002032225 A2 20020425 (200251)* EN
PΙ
                                              32p
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ
            LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT/RO RU SD
            SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW
     AU 2002011792 A 20020429 (200255)
    WO 2002032225 A2 WO 2001-US32415 20011018; AU 2002011792 A AV 2002-11792
ADT
     20011018
    AU 2002011792 A Based, on WO 200232225
FDT
PRAI US 2000-691197
                     2000/10/19
     WO 200232225 A UPAB: 20020812
     NOVELTY - A natural or engineered tissue or organ
     other than a blood vessel is vitrified by:
          (i) immersing it in a series of solutions having increasing
     concentrations of cryoprotectant to achieve a cryoprofectant concentration
     for vitrification;
          (ii) rapidly cooling to between -80 deg. C and /the glass transition
     temperature (Tg); and
          (iii) further cooling to below Tg.
          DETAILED DESCRIPTION - Vitrification of a natural or engineered
     tissue or organ other than a blood vessel comprises:
          (i) immersing the tissue (3) or organ in a series
     of solutions having increasing concentrations of cryoprotectant
     and each having a temperature above -15 deg. C;
          (ii) cooling the tissue of organ at 2.5-100 deg.
     C per minute from a temperature above -15 deg. C to between -80 deg. C and
     the Tg; and
          (iii) further cooling at average rate less than 30 deg. C per minute
     from between -80 deg. C and the Tg to below the Tg to vitrify the
     tissue or organ.
          An INDEPENDENT CLAIM is also included for a method for removing a
     tissue or organ other than a blood from vitrification in
     a solution containing cryoprotectant by:
          (a) warming the vitrified tissue or organ in a
     solution containing cryoprotectant at an average rate of 20-40
     deg. C per minute to between -80 deg. C and the Tg;
          (b) further warming in the solution at an average rate of
     200-300 deg. C per minute to a temperature above -75 deg. C; and
          (c) immersing the tissue or organ in a series of
     solutions having decreasing concentrations of cryoprotectant.
          USE - For vitrifying a natural or engineered tissue or
     organ other than a blood vessel, particularly musculoskeletal
     tissue, cartilage, menisci, muscles, ligaments, tendons, skin,
     cardiovascular tissue, heart valves, myocardium, periodontal
     tissue, qlandular tissue, islets of Lange, cornea,
     ureter, urethra, pancreas, bladder, kidney, breast, liver, intestine or
          ADVANTAGE - The vitrification method results in a greater number or
     percentage of viable cells in a tissue or
```

organ sample, compared to conventional cryopreservation

techniques. It results in **tissue** or **organ** samples having at least 50% viable **cells**.

DESCRIPTION OF DRAWING(S) - The figure shows a perfusion system that can be used in the invention.

Tissue 3

Dwg.1/4

TI Vitrification of natural or engineered tissue or organ other than blood vessel involves immersing tissue in cryoprotectant solutions with increasing concentrations and cooling to below glass transition temperature.

AB WO 200232225 UPAB: 20020812

NOVELTY - A natural or engineered tissue or organ other than a blood vessel is vitrified by:

- (i) immersing it in a series of **solutions** having increasing concentrations of cryoprotectant to achieve a cryoprotectant concentration for vitrification;
- (ii) rapidly cooling to between -80 deg. C. . . glass transition temperature (Tg); and

(iii) further cooling to below Tg.

DETAILED DESCRIPTION - Vitrification of a natural or engineered tissue or organ other than a blood vessel comprises:

- (i) immersing the **tissue** (3) or **organ** in a series of **solutions** having increasing concentrations of cryoprotectant and each having a temperature above -15 deg. C;
- (ii) cooling the **tissue** of **organ** at 2.5-100 deg. C per minute from a temperature above -15 deg. C to between -80 deg. C and the. . . 30 deg. C per minute from between -80 deg. C and the Tg to below the Tg to vitrify the **tissue** or **organ**.

An INDEPENDENT CLAIM is also included for a method for removing a tissue or organ other than a blood from vitrification in a solution containing cryoprotectant by:

- (a) warming the vitrified **tissue** or **organ** in a **solution** containing cryoprotectant at an average rate of 20-40 deg. C per minute to between -80 deg. C and the Tg;
- (b) further warming in the **solution** at an average rate of 200-300 deg. C per minute to a temperature above -75 deg. C; and
- (c) immersing the tissue or organ in a series of solutions having decreasing concentrations of cryoprotectant.

USE - For vitrifying a natural or engineered tissue or organ other than a blood vessel, particularly musculoskeletal tissue, cartilage, menisci, muscles, ligaments, tendons, skin, cardiovascular tissue, heart valves, myocardium, periodontal tissue, glandular tissue, islets of Lange, cornea, ureter, urethra, pancreas, bladder, kidney, breast, liver, intestine or heart.

ADVANTAGE - The vitrification method results in a greater number or percentage of viable **cells** in a **tissue** or **organ** sample, compared to conventional cryopreservation techniques. It results in **tissue** or **organ** samples having at least 50% viable **cells**.

 $\overline{\text{DESCRIPTION}}$ OF DRAWING(S) - The figure shows a perfusion system that can be used in the invention.

Tissue 3

Dwg.1/4

TECH UPTX: 20020812

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Method: Cooling the tissue or organ from above -15 degreesC to between -80 degreesC and the Tg is performed at an average rate of at least 10 degreesC (preferably 30-60 degreesC) per minute. Cooling the tissue or organ from between -80 degreesC and the Tg to below the Tg is performed at an average rate less than 10 degreesC per minute. The immersion step (i) comprises immersing the tissue or organ in a cryoprotectant-free solution; immersing the tissue or organ in solution(s) containing

cryoprotectant at a concentration less than the concentration sufficient for the vitrification; and immersing the **tissue** or **organ** in a **solution** containing cryoprotectant at the concentration sufficient for vitrification. In each immersion step, the **tissue** or **organ** is immersed in the **solution** for at least 10 minutes. The increasing and decreasing concentrations of cryoprotectant are (a) 5-50%, 15-35%, 40-60%, and 65-85%; or (b) 40-60%, 30-45%, 15-35%, 5-20%, 2.5-10%, and 0%.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Components: The cryoprotectant solution comprises acetamide, agarose, alginate, alanine, albumin, ammonium acetate, anti-freeze proteins, butanediol, chondroitin sulfate, chloroform, choline, cyclohexanediols, dextrans, diethylene glycol, dimethyl acetamide, dimethyl formamide, dimethyl sulfoxide, erythritol, ethanol, ethylene glycol, ethylene glycol monomethyl ether, formamide, glucose, glycerol, glycerophosphate, glyceryl monoacetate, glycine, glycoproteins, hydroxyethyl starch, inositol, lactose, magnesium chloride, magnesium sulfate, maltose, mannitol, mannose, methanol, methoxy propanediol, methyl acetamide, methyl formamide, methyl ureas, methyl glucose, methyl glycerol, phenol, pluronic polyols, polyethylene glycol, polyvinyl pyrrolidone, proline, 1,2-propanediol, pyridine N-oxide, raffinose, ribose, serine, sodium bromide, sodium chloride, . . . and/or xylose. It preferably comprises (in weight per volume) dimethyl sulfoxide (20-30%), formamide (10-20%) and 1,2-propanediol (10-20%) in a vehicle solution. Each solution comprises an osmotic buffering agent, preferably mannitol.

TT: VITREOUS NATURAL ENGINEERING TISSUE ORGAN BLOOD
VESSEL IMMERSE TISSUE SOLUTION INCREASE
CONCENTRATE COOLING BELOW GLASS TRANSITION TEMPERATURE.

=>

- L7 ANSWER 23 OF 37 CA COPYRIGHT 2003 ACS
- AN 121:252473 CA
- Effect of polyols on the post-thawing motility of pellet-frozen ram ΤI spermatozoa
- Molinia, F. C.; Evans, G.; Maxwell, W. M. C. ΑU
- Department Animal Science, University Sydney, Sydney, NSW 2006, Australia CS
- SO Theriogenology (1994), 42(1), 15-23 CODEN: THGNBO; ISSN: 0093-691X

Tris-glucose-egg yolk diluents.

- DTJournal
- English LΑ
- The cryoprotective effects of polyols in the absence and presence of AΒ glycerol in Tris-glucose-egg yolk based diluents on the post-thawing motility and acrosome integrity of pellet-frozen ram spermatozoa were examd. Incorporation of adonitol or xylitol (low mol. wt. polyols; LMWPs) in diluents improved motility of spermatozoa in the absence of glycerol with max. motility at 0.3 M (26.9 vs. 13.3% mean motile spermatozoa). Five concns. of adonitol (0, 150, 300, 450, 600 mM) were examd. in diluents contg. 5 concns. of glycerol (0, 1.5, 3.0, 4.5, 6.0% vol./vol.). There was an additive effect of incorporation of 1.5% vol./vol. glycerol with up to 450 mM adonitol, but at higher levels of glycerol the incorporation of adonitol was detrimental to motility. The acrosome integrity of spermatozoa in diluents contg. 0, 150 and 300 mM adonitol was superior to those contg. 450 and 600 mM adonitol (46.1 vs. 35.1% mean intact acrosomes). Among the high mol. wt. polyols (HMWPs) examd., better recovery of spermatozoa was obtained in diluents contg. sorbitol than mannitol or inositol. Sorbitol or mannitol (300 mM) improved the motility of spermatozoa in diluents without glycerol, but the incorporation of HMWPs was detrimental in diluents contg. glycerol. All five polyols were examd. in isotonic diluents contg. 360:0, 300:55, 240:110, 180:165, 120:220mM (Tris:polyol; 360 mosmol) and 6.0% vol./vol. glycerol. There was a linear decrease in motility and acrosome integrity of spermatozoa with increasing polyol concn. in the diluent except for inositol, which was not detrimental. The authors conclude that the polyols examd. have a cryoprotective effect on pellet-frozen ram spermatozoa except for inositol. However, in the authors study, no combination of polyols and glycerol was superior in terms of post-thawing motility and acrosome integrity of spermatozoa to 6.0% vol./vol. glycerol alone in

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS

RN 9041-07-0 REGISTRY

CN Decaglycerol (9CI) (CA INDEX NAME)

OTHER NAMES:

=>

CN Decaglycerin

CN Polyglycerin 10

DR 83689-42-3, 26085-10-9, 34322-27-5

MF C30 H62 O21

CI IDS, COM, MAN

LC STN Files: BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CHEMLIST, IFICDB, IFIPAT, IFIUDB, NIOSHTIC, TOXCENTER, USPATFULL

Other Sources: EINECS**, NDSL**, TSCA**

(**Enter CHEMLIST File for up-to-date regulatory information)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

127 REFERENCES IN FILE CA (1962 TO DATE)

63 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

127 REFERENCES IN FILE CAPLUS (1962 TO DATE)

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1998-076784 [07]
                        WPIDS
AN
     1998-593980 [50]
CR
    C1998-025629
DNC
     Solution for preservation of biological materials - comprises
ΤI
     two neutral solutes, especially raffinose and tri methylamine oxide.
     B04 D16 D22 E19 E37 P31
DC
     FERGUSON, A B; WIGGINS, P M; WATSON, J D
IN
     (BIOS-N) BIOSTORE NEW ZEALAND LTD; (BIOS-N) BIOSTORE NZ LTD; (BIOS-N)
PA
     BIOSTORE NEW ZEALAND
CYC
                                                     A01N001-00
                   A1 19971218 (199807)* EN
                                              53p
    (WO 9747192)
PΙ
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            SE SZ UG
         W: AL AM AT AU AZ BB BG BR BY CA CH CN CZ DE DK EE ES FI GB GE HU IL
            IS JP KE KG KP KR KZ LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL
            PT RO RU SD SE SG SI SK TJ TM TR TT UA UG UZ VN
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     AII 9661412
                                                     A61K000-00
                                              79p
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     ZA 9710452
                   Α
                                                     A01N001-02
                   A 19990309 (199917)#
     US 5879875
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                   A 20000526 (200033)
     NZ 333030
                                                     A01N001-00
                   A1 20000719 (200036)
                                         EN
     EP 1018866
         R: CH DE FR GB LI SE
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                                              50p
     JP 2000512625 W
                     20000926 (200051)
                                                     A01N001-00
                   B 20001012 (200055)
     AU 725247
                                                     A61K035-28
     AU 2001010037 A 20010315 (200121)#
                                                     A61K035-28
     AU 742402
                   B 20020103 (200209)#
ADT WO 9747192 A1 WO 1996-NZ57 19960614; AU 9661412 A AU 1996-61412 19960614,
     WO 1996-NZ57 19960614; ZA 9710452 A ZA 1997-10452 19971120; US 5879875 A
     US 1996-662244 19960614; NZ 333030 A NZ 1996-333030 19960614, WO
1996-NZ57
     19960614; EP 1018866 A1 EP 1996-918938 19960614, WO 1996-NZ57 19960614;
JP
     2000512625 W WO 1996-NZ57 19960614, JP 1997-541244 19960614; AU 725247 B
     AU 1996-61412 19960614, WO 1996-NZ57 19960614; AU 2001010037 A Div ex AU
     1996-61412 19960614, AU 2001-10037 20010103; AU 742402 B Div ex AU
     1996-61412 19960614, AU 2001-10037 20010103
FDT AU 9661412 A Based on WO 9747192; NZ 333030 A Based on WO 9747192; EP
     1018866 A1 Based on WO 9747192; JP 2000512625 W Based on WO 9747192; AU
     725247 B Previous Publ. AU 9661412, Based on WO 9747192; AU 2001010037 A
     Div ex AU 725247; AU 742402 B Previous Publ. AU 200110037, Div ex AU
     725247
                                                  19971120; US 1996-662244
                      19960614; ZA 1997-10452
PRAI WO 1996-NZ57
     19960614; AU 2001-10037
                                20010103
     ICM A01N001-00; A01N001-02; A01N003-00; A61K000-00; A61K035-28
IC
          A61B000-00; A61K031-00; A61K035-12; A61K035-14; A61K035-34;
     ICS
          A61K035-36; A61K035-60; A61K035-78; A61P035-02; C12N001-04;
          C12N005-00; C12N005-06
          9747192 A UPAB: 20020208
AR
     WO
     The following are claimed: (1) a solution for preservation of
     biological materials: (A) comprising: (a) a first neutral solute with a
     molecular weight of at least 335 and a solubility in water of at least
0.3
     M, and (b) a second neutral solute with a molecular weight < 200 and with
     both hydrophilic and hydrophobic moieties; or (B) which is isotonic with
     the biological materials and is free of univalent oxyanions and iodide,
     and (2) preservation of biological materials by: (a) pretreating the
     biological material with a solution which includes sodium
     butyrate, and (b) contacting the biological material with a preservative
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solution.

The **solution** (A) is free of univalent oxyanions and iodide, and also comprises at least 1 ion from a protein-stabilising end of the Hofmeister series. The first solute is selected from disaccharides and trisaccharides, especially raffinose, trehalose, sucrose, **lactose** and their analogues. The second neutral solute is selected from trimethylamine oxide, betaine, taurine, sarcosine, glucose, mannose, fructose, ribose, galactose, sorbitol, **mannitol**, inositol and their analogues. The **solution** may also comprise sodium sulphate. It may also comprise calcium, which is present as calcium sulphate at a concentration of 1.5-2.0 mM. The **solution** is in a concentrated form, especially in the form of a solid. Components (a) and (b) are typically present at a ratio of 1.4-1.8:1.

USE - The **solutions** may be used for preservation of materials such as organs, tissues and cells from mammals, marine organisms

and plants. They may be used e.g. in treatment of leukaemia. In this case, bone marrow is removed from a patient and contacted with the solution for at least 3 days, in order to purge the bone marrow of leukaemic cells. The purged bone marrow is then returned to the patient.

ADVANTAGE - The solutions are of low toxicity, resulting in fewer side effects when biological materials, such as transplant organs, are returned to a patient.

Dwg.0/20

FS CPI GMPI

FA AB; DCN

MC CPI: B04-C02X; B10-A07; B14-H01A; B14-N17B; D05-H01; D09-A01; E07-A02A; E07-A02D; E07-A02H; E10-A03

- L28 ANSWER 30 OF 31 CA COPYRIGHT 2002 ACS
- AN 98:213494 CA
- TI Identification of new cryoprotective agents for cultured mammalian cells
- AU Klebe, Robert J.; Mancuso, Melodee G.
- CS Dep. Anat., Univ. Texas Health Sci. Cent., San Antonio, TX, 78284, USA
- SO In Vitro (1983), 91(3, pt. 1), 167-70 CODEN: ITCSAF; ISSN: 0073-5655
- DT Journal
- LA English
- AB Thirty-one compds. were identified that act as cryoprotective agents for cultured mammalian (CHO) cells. Eight compds. were comparable to DMSO in cryoprotective effectiveness. Many of the cryoprotective compds. studied also (1) promote cell fusion and (2) induce cell differentiation in erythroleukemia and other cell systems. Thus, previously unrecognized effects on the differentiated state of cells may occur when cells are treated with cryoprotective agents.

ANSWER 7 OF 30 MEDLINE

ΑN 1998353576 MEDLINE

98353576 PubMed ID: 9687336 DN

- ΤI The physical state of mannitol after freeze-drying: effects of mannitol concentration, freezing rate, and a noncrystallizing cosolute.
- ΑU Kim A I; Akers M J; Nail S L
- CS Department of Industrial and Physical Pharmacy, Purdue University, West Lafayette, Indiana 47907, USA.
- SO JOURNAL OF PHARMACEUTICAL SCIENCES, (1998 Aug) 87 (8) 931-5. Journal code: 2985195R. ISSN: 0022-3549.
- CY United States
- DТ Journal; Article; (JOURNAL ARTICLE)
- English LΑ
- FS Priority Journals
- ΕM 199809
- Entered STN: 19980910 ED Last Updated on STN: 19980910 Entered Medline: 19980903
- AB The objectives of this study were to (1) measure the effects of freezing rate and mannitol concentration on the physical state of freeze-dried mannitol when mannitol is present as a single component, (2) determine the relative concentration threshold above

which crystalline mannitol can be observed by X-ray powder diffraction in the freeze-dried solid when a variety of noncrystallizing solutes are included in the formulation, and (3) measure the glass transition temperature of amorphous mannitol and to determine the degree to which the glass transition temperature of freeze-dried solids consisting of mannitol and a disaccharide is predicted by the Gordon-Taylor equation. Both freezing rate and mannitol concentration influence the crystal form of mannitol in the freeze-dried solid when mannitol is present as a single component. Slow freezing of 10% (w/v) mannitol produces a mixture of the alpha and beta polymorphs, whereas fast freezing of the same solution produces the delta form. Fast freezing of 5% (w/v)mannitol results primarily in the beta form. The threshold concentration above which crystalline mannitol is detected in the freeze-dried solid by X-ray diffraction is consistently about 30% (w/w) when a second, noncrystallizing solute is present, regardless of

the

nature of the second component. The glass transition temperature of amorphous mannitol measured from the quench-cooled melt is approximately 13 degreesC. Accordingly, mannitol is an effective plasticizer of freeze-dried solids when the mannitol remains amorphous. Glass transition temperatures of mixtures of mannitol and the disaccharides sucrose, maltose, trehalose, and lactose are well predicted by the Gordon-Taylor equation with values of k in the range of 3 to 4.

30,28,25,13,10,8

9041-07-0 9002-69-5 X

L9 ANSWER 184 OF 268 WPIDS (C) 2002 THOMSON DERWENT AN 1989-162877 [22] WPIDS DNC C1989-072372 New blood-preservative liq. compsn. - contg. haemolysis preventive of ΤI mono carboxylic ester and other blood-preservative liq. ingredients, esp. for red blood corpuscles. DC B05 D22 E19 PA (KAOS) KAO CORP; (TERU) TERUMO CORP CYC PΙ JP 01106826 A 19890424 (198922)* q8 ADT JP 01106826 A JP 1987-263695 19871021 PRAI JP 1987-263695 19871021 IC A61K035-14 AB JP 01106826 A UPAB: 19930923 New blood-preservative liq. compsn. contains a haemolysis preventive of monocarboxylic ester of formula RCOOR' (I) and other blood-preservative liq. ingredients. (R and R' (individually) are 3C or higher chain hydrocarbon with R + R' = 11-30). Blood-preservative ligs. are e.g. anticoagulants and at least one of sodium citrate, citric acid, glucose, monosodium phosphate, adenine, sodium chloride, mannitol, maltose, multitol, sorbitol, sucrose, and lactose. Compsn. is prepd. by blending (1) with base liq. of ACd, CPD, CPDA-1, CPDA-2, SAG and SAG solns. contg. mannitol, maltose, maltose,

multitol, sorbitol, martose, maltose, multitole, multitol, sorbitol, sucrose, and lactose. Either or both of R and R' are pref. branched. Final concn. of ester is 100 M - 10mM; and concn. is 400microM - 4 mM for branched R and/or R'.

USE/ADVANTAGE - Compsn. with high physiological safety, has good

protective action esp. on red blood corpuscles. Effect is durable. 0/0

FS